

- 1 5. The nucleic acid molecule of Claim 2, which has a sequence as shown in
2 Figure 1 (SEQ ID NO:1) from nucleotide number 46 to nucleotide number 550.
- 1 6. The nucleic acid molecule of Claim 2, which has a sequence as shown in
2 Figure 2 (SEQ ID NO:2) from nucleotide number 46 to nucleotide number 550.
- 1 7. The nucleic acid molecule of Claim 1 which is detectably labeled.
- 1 8. A cloning vector, which comprises the DNA molecule of Claim 1.
- 1 9. An expression vector, which comprises the nucleic acid molecule of Claim
2 2, operatively associated with an expression control sequence.
- 1 10. The expression vector of Claim 9, wherein said expression control
2 sequence is selected from the group consisting of the cytomegalovirus hCMV
3 immediate early gene, the early or late promoters of SV40 or adenovirus, the lac
4 system, the trp system, the TAC system, the TRC system, the major operator and
5 promoter regions of phage λ , the control regions of fd coat protein, the promoter
6 for 3-phosphoglycerate kinase, the promoters of acid phosphatase, and the
7 promoters of the yeast α -mating factors.
- 1 11. A probe capable of screening for a nucleic acid encoding an ob polypeptide
2 in alternate species, which probe is a labeled DNA molecule of Claim 1.
- 1 12. A unicellular host transfected with a cloning vector of Claim 8.
- 1 13. A unicellular host transfected with an expression vector of Claim 9.
- 1 14. The unicellular host of Claim 13 wherein the unicellular host is selected
2 from the group consisting of *E. coli*, *Pseudomonas*, *Bacillus*, *Streptomyces*, *Pichia*

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3 yeasts, CHO, R1.1, B-W, L-M, COS 1, COS 7, BSC1, BSC40, and BMT10 cells,
4 plant cells, insect cells, and human cells in tissue culture.

1 15. An oligonucleotide primer for amplifying human genomic DNA encoding
2 an ob polypeptide.

1 16. The oligonucleotide of Claim 15, which is selected from the group
2 consisting of

3 HOB 1gF 5'-CCCAAGAAGCCCATCCTG-3' (SEQ ID NO:26)

4 HOB 1gR 5'-GACTATCTGGGTCCAGTGCC-3' (SEQ ID NO:27)

5 HOB 2gF 5'-CCACATGCTGAGCACTTGTT-3' (SEQ ID NO:28)

6 HOB 2gR 5'-CTTCAATCCTGGAGATACCTGG-3' (SEQ ID NO:29).

1 17. An ob polypeptide, which polypeptide is encoded by the DNA molecule of
2 Claim 1.

1 18. An ob polypeptide, which polypeptide is characterized by having about 145
2 to about 167 amino acid residues, being expressed predominantly by adipocytes,
3 and being capable of inducing a reduction of body weight in an animal.

1 19. The ob polypeptide of Claim 18 which has the amino acid sequence shown
2 in Figure 1 (SEQ ID NO:2) or Figure 5 (SEQ ID NO:5).

1 20. The ob polypeptide of Claim 19 which has the amino acid sequence shown
2 in Figure 3 (SEQ ID NO:4) or Figure 6 (SEQ ID NO:6).

1 21. An immunogenic fragment of an ob polypeptide, which polypeptide is
2 characterized by having about 160 amino acid residues, being expressed
3 predominantly by adipocytes, and being capable of inducing a reduction of body
4 weight in an animal.

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1 22. The immunogenic fragment of an ob polypeptide of Claim 21, which is
2 selected from the group consisting of

3 Val-Pro-Ile-Gln-Lys-Val-Gln-Asp-Asp-Thr-Lys-Thr-Leu-Ile-Lys-Thr (SEQ
4 ID NO:18);

5 Leu-His-Pro-Ile-Leu-Ser-Leu-Ser-Lys-Met-Asp-Gln-Thr-Leu-Ala (SEQ ID
6 NO:19);

7 Ser-Lys-Ser-Cys-Ser-Leu-Pro-Gln-Thr-Ser-Gly-Leu-Gln-Lys-Pro-Glu-Ser-
8 Leu-Asp (SEQ ID NO:20); and

9 Ser-Arg-Leu-Gln-Gly-Ser-Leu-Gln-Asp-Ile-Leu-Gln-Gln-Leu-Asp-Val-Ser-
10 Pro-Glu-Cys (SEQ ID NO:21).

1 23. A method for preparing an ob polypeptide comprising:

2 A. culturing a unicellular host of Claim 12 or 13 under conditions that
3 provide for expression of the ob polypeptide; and

4 B. recovering the expressed ob polypeptide.

5 24. The method according to Claim 23 wherein the host cell is a bacterium.

1 25. The method according to Claim 23, wherein the host cell is a yeast.

1 26. The method according to Claim 23, further comprising:

2 C. chromatographing the polypeptide on a Ni-chelation column; and

3 D. purifying the polypeptide by gel filtration.

1 27. The method according to Claim 26, further comprising after step C and
2 before step D chromatographing the ob polypeptide on a strong cation exchanger
3 column.

1 28. An antibody to the ob polypeptide of Claim 17.

1 29. An antibody to the ob polypeptide of Claim 18.

- 1 30. A method for preparing an antibody to an ob polypeptide, comprising:
2 A. conjugating the immunogenic fragment of an ob polypeptide of
3 Claim 19 to a carrier protein;
4 B. immunizing a host animal with the ob polypeptide fragment-carrier
5 protein conjugate of step A admixed with an adjuvant; and
6 C. obtaining antibody from the immunized host animal.
- 1 31. An antibody to an ob polypeptide prepared according to a method
2 comprising:
3 A. conjugating an immunogenic fragment of an ob polypeptide of
4 Claim 19 to a carrier protein;
5 B. immunizing a host animal with the ob polypeptide fragment-carrier
6 protein conjugate of step A admixed with an adjuvant; and
7 C. obtaining antibody from the immunized host animal.
- 1 32. The antibody of Claim 28, 29, or 31 comprising a polyclonal antibody.
- 1 33. The antibody of Claim 28, 29, or 30 comprising a monoclonal antibody.
- 1 34. An immortal cell line that produces a monoclonal antibody according to
2 Claim 33.
- 1 35. The antibody of Claim 28, 29, or 31 labeled with a detectable label.
- 1 36. The antibody of Claim 35 wherein the label is selected from the group
2 consisting of enzymes, chemicals which fluoresce, and radioactive elements.
- 1 37. A method for measuring the presence of an ob polypeptide in a sample,
2 comprising:
3 A. contacting a sample suspected of containing an ob polypeptide with
4 an antibody that binds to the ob polypeptide under conditions which allow for the

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1 43. A method for evaluating the level of ob polypeptide in a biological sample
2 comprising

3 A. detecting the formation of reaction complexes in a biological sample
4 according to the method of Claim 30; and

5 B. evaluating the amount of reaction complexes formed, which amount
6 of reaction complexes corresponds to the level of ob polypeptide in the biological
7 sample.

1 44. A method for detecting or diagnosing the presence of a disease associated
2 with elevated or decreased levels of ob polypeptide in a mammalian subject
3 comprising:

4 A. evaluating the level of ob polypeptide in a biological sample from a
5 mammalian subject according to Claim 43; and

6 B. comparing the level detected in step (A) to a level of ob polypeptide
7 present in normals or in the subject at an earlier time;
8 in which an increase in the level of ob polypeptide as compared to normal levels
9 indicates a disease associated with elevated levels of ob polypeptide, and decreased
10 level of ob polypeptide as compared to normal levels indicates a disease associated
11 with decreased levels of ob polypeptide.

1 45. A method for monitoring a therapeutic treatment of a disease associated
2 with elevated or decreased levels of ob polypeptide in a mammalian subject
3 comprising evaluating the levels of ob polypeptide in a series of biological samples
4 obtained at different time points from a mammalian subject undergoing a
5 therapeutic treatment for a disease associated with elevated or decreased levels of
6 ob polypeptide according to the method of Claim 43.

1 46. The method according to Claim 44 or 45, wherein the disease associated
2 with elevated levels of ob polypeptide is selected from the group consisting of
3 AIDS, cachexia, cancer, and anorexia nervosa.

1 52. A pharmaceutical composition for reducing body weight of an animal
2 comprising the ob polypeptide of Claim 18 and a pharmaceutically acceptable
3 carrier.

1 53. A method for reducing the body weight of an animal comprising
2 administering an amount of a pharmaceutical composition of Claim 52 effective to
3 reduce the body weight of an animal to an animal believed to be in need of
4 decreased body weight.

1 54. The method according to Claim 53 wherein the animal is a human, and the
2 ob polypeptide is human ob polypeptide.

1 55. A method for reducing the body weight of a mammal comprising increasing
2 the expression of a protein encoded by the nucleic acid of Claim 2.

1 56. A pharmaceutical composition for increasing the body weight of an animal
2 comprising an antagonist of an ob polypeptide.

1 57. The pharmaceutical composition of Claim 56, wherein the antagonist is
2 selected from the group consisting of an antibody that binds to and neutralizes the
3 activity of ob polypeptide, a fragment of the ob polypeptide that binds to but does
4 not activate the ob receptor, and a small molecule antagonist of the ob
5 polypeptide.

1 58. A method for increasing the body weight of an animal comprising
2 administering an amount of the pharmaceutical composition of Claim 56 effective
3 to cause an increase in body weight to an animal believed to be in need of
4 increased body weight.

add B.

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